

```
chain nodes :
    16 24 26 37
ring nodes :
     1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 18 19 20 21 22 23 27 28 29 30
     31 32
ring/chain nodes :
     38 39
chain bonds:
     7-10 16-37 26-27
ring/chain bonds :
     8-38 8-39
ring bonds :
     1-2 1-6 2-3 3-4 4-5 5-6 5-7 6-9 7-8 8-9 10-11 10-15 11-12 12-13 13-14 14-15 18-19 18-23 19-20 20-21 21-22 22-23 27-28 27-32 28-29 29-30 30-31 31-32
exact/norm bonds :
     7-8 8-38 8-39 16-37 26-27
exact bonds:
     5-7 6-9 7-10 8-9
normalized bonds :
     1-2 1-6 2-3 3-4 4-5 5-6 10-11 10-15 11-12 12-13 13-14 14-15 18-19 18-23 19-20 20-21 21-22 22-23 27-28 27-32 28-29 29-30 30-31 31-32
isolated ring systems:
     containing 1 : 10 : 18 : 27 :
G1:[*1],[*2]
Match level:
     1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom 11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 16:CLASS 17:CLASS 18:Atom 19:Atom 20:Atom 21:Atom 22:Atom 23:Atom 24:CLASS 25:CLASS 26:CLASS 27:Atom 28:Atom 29:Atom
```

30:Atom 31:Atom 32:Atom 37:CLASS 38:CLASS 39:CLASS

* * *	* *	* *	* *	* Welcome to STN International * * * * * * * * *
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NEWS	3	SEP	01	New pricing for the Save Answers for SciFinder Wizard within
				STN Express with Discover!
NEWS	4	OCT	28	KOREAPAT now available on STN
NEWS	5	VОИ	18	Current-awareness alerts, saved answer sets, and current
				search transcripts to be affected by CERAB, COMPUAB, ELCOM,
				and SOLIDSTATE reloads
NEWS	6	NOV	30	PHAR reloaded with additional data
NEWS		DEC	01	LISA now available on STN
NEWS	8	DEC	09	12 databases to be removed from STN on December 31, 2004
NEWS	9	DEC	15	MEDLINE update schedule for December 2004
NEWS	EXP	RESS		FOBER 29 CURRENT WINDOWS VERSION IS V7.01A, CURRENT
				CINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),
				CURRENT DISCOVER FILE IS DATED 11 AUGUST 2004
NEWS	HOU	RS	STI	N Operating Hours Plus Help Desk Availability
NEWS	INT	ER	Gei	neral Internet Information
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=> file reg
COST IN U.S. DOLLARS
SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST
0.21
0.21

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STRUCTURE FILE UPDATES: 14 DEC 2004 HIGHEST RN 797749-23-6 DICTIONARY FILE UPDATES: 14 DEC 2004 HIGHEST RN 797749-23-6

TSCA INFORMATION NOW CURRENT THROUGH MAY 21, 2004

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Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at: http://www.cas.org/ONLINE/DBSS/registryss.html

=>

L1 STRUCTURE UPLOADED

=> 11

L1 IS NOT A RECOGNIZED COMMAND

The previous command name entered was not recognized by the system. For a list of commands available to you in the current file, enter "HELP COMMANDS">HELP COMMANDS" at an arrow prompt (=>).

=> s l1

SAMPLE SEARCH INITIATED 12:59:47 FILE 'REGISTRY' SAMPLE SCREEN SEARCH COMPLETED - 44 TO ITERATE

100.0% PROCESSED

44 ITERATIONS

1 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**

PROJECTED ITERATIONS:

483 TO 1277

PROJECTED ANSWERS:

1 TO 8

L2

1 SEA SSS SAM L1

=> s 11 full

THE ESTIMATED SEARCH COST FOR FILE 'REGISTRY' IS 155.00 U.S. DOLLARS DO YOU WANT TO CONTINUE WITH THIS REQUEST? (Y)/N or END: Y FULL SEARCH INITIATED 12:59:51 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 904 TO ITERATE

100.0% PROCESSED

904 ITERATIONS

24 ANSWERS

SEARCH TIME: 00.00.01

 Γ 3

h

24 SEA SSS FUL L1

=> file hcaplus

COST IN U.S. DOLLARS

SINCE FILE TOTAL

ENTRY SESSION 157.52 157.73

FULL ESTIMATED COST

FILE 'HCAPLUS' ENTERED AT 12:59:53 ON 16 DEC 2004 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

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FILE COVERS 1907 - 16 Dec 2004 VOL 141 ISS 25 FILE LAST UPDATED: 15 Dec 2004 (20041215/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s 13

L4

4 L3

=> d 14, ibib abs hitstr, 1-4

ANSWER 1 OF 4 HCAPLUS COPYRIGHT 2004 ACS on STN L4

ACCESSION NUMBER:

2003:719300 HCAPLUS

DOCUMENT NUMBER:

139:240389

TITLE:

Antidepressant

INVENTOR(S): PATENT ASSIGNEE(S): Ohkawa, Shigenori; Miyamoto, Masaomi Takeda Chemical Industries, Ltd., Japan

SOURCE:

PCT Int. Appl., 95 pp.

DOCUMENT TYPE:

CODEN: PIXXD2

LANGUAGE:

Patent Japanese

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

	PAT	rent :	NO.			KIN	D	DATE			APPL					D.	ATE	
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			CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FΙ,	GB,	GD,	GE,	GH,
			GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	ΚG,	KR,	ΚZ,	LC,	LK,	LR,	LS,
			LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NO,	NZ,	OM,	PH,	PL,
			PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	ТJ,	TM,	TN,	TR,	TT,	TZ,	UA,
	UG, US, U					VC,	VN,	YU,	ZA,	ZM,	zw							
	RW: GH, GM, K					LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	ΑZ,	BY,
	KG, KZ, M					RU,	ТJ,	TM,	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	ES,
			FI,	FR,	GB,	GR,	HU,	ΙE,	IT,	LU,	MC,	NL,	PT,	SE,	SI,	SK,	TR,	BF,
			ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,	TD,	$\mathbf{T}G$	
	JP	2004	0835	56		A2		2004	0318		JP 2	003-	5250	3		2	0030	228
	EP	1481	679			A1		2004	1201		EP 2	003-	7071	69		2	0030	228
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PRIO	RITY	APP	LN.	INFO	.:						JP 2	002-	5577	1	1	A 2	0020	301
											JP 2	002-	1954	34	i	A 2	0020	704
											WO 2	003-	JP22	93	1	w 2	0030	228
OTHE	3 50	URCE	(S):			MAR	PΆΨ	139:	24038	39								

OTHER SOURCE(S):

MARPAT 139:240389

GI

$$\mathbf{N} = \mathbf{N} =$$

A PKB (Akt) activator contg. a compd. represented by the formula (I) [wherein R1 and R2 each represents hydrogen, a hydrocarbon group, or a heterocyclic group or R1 and R2 form a ring in cooperation with the adjacent carbon atom; R3 represents hydrogen, a hydrocarbon group, or a heterocyclic group; W represents a group represented by the formula (II) (-N(R4)(R5)) or (-XR4c) (wherein ring A represents an optionally substituted benzene ring; ring B represents an optionally substituted 5-to 7-membered nitrogenous heterocycle; R4 represents either an arom.-group-substituted aliph. hydrocarbon group which may have other substituent(s) or an acyl contg. an arom. group; R5 represents hydrogen, C1-6 alkyl, or acyl; R4c represents an arom. group, aliph. hydrocarbon group, or acyl; and X represents oxygen or sulfur); Y represents oxygen, sulfur, or NH; and ring C represents an optionally substituted benzene ring], a salt of the compd., or a prodrug of either. Also provided is a use of the activator in or as a preventive/therapeutic agent for depressive psychoses, anxiety disorders, affective psychoses, or PTSD.

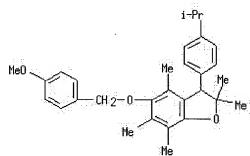
IT 216989-18-3

RN

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(benzofuran analogs as protein kinase B activators and antidepressants) 216989-18-3 HCAPLUS

CN Benzofuran, 2,3-dihydro-5-[(4-methoxyphenyl)methoxy]-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT:

THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 2 OF 4 HCAPLUS COPYRIGHT 2004 ACS on STN

3

Full Text diekelde

ACCESSION NUMBER: 2002:275980 HCAPLUS

DOCUMENT NUMBER: 136:309840

TITLE: Preparation of heterocyclic compounds as promoters for

the proliferation and differentiation of stem cells

and neuron precursor cells

INVENTOR(S): Okawa, Shigenori; Miyamoto, Masaomi; Okura, Masahiro

PATENT ASSIGNEE(S): Takeda Chemical Industries, Ltd., Japan

SOURCE: PCT Int. Appl., 182 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent
LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PAT	rent	NO.			KIN	D :	DATE			APPL	ICAT	ION	NO.		D	ATE	
	2002	0200	-		A1	-	2002	0411			001		20			0011	
WO	2002	0200	30		AI		2002	0411		WO 2	00T-	<u> 1587</u>	39		2	0011	004
	W:	ΑE,	AG,	AL,	AM,	ΑT,	AU,	AZ,	BA,	BB,	ВG,	BR,	BY,	BZ,	CA,	CH,	CN,
		co,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	ΕĒ,	ES,	FI,	GB,	GD,	GE,	GH,
		GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KR,	ΚZ,	LC,	LK,	LR,	LS,
		LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NO,	NZ,	PH,	PL,	PT,

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RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US,
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         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
             BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
    AU 2001092350
                          Α5
                                20020415
                                            AU 2001-92350
                                                                    20011004
     JP 2002348239
                                            JP 2001-308530
                                                                    20011004
                          A2
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                                            CA 2001-2424870
    CA 2424870
                          AA
                                20030404
                                                                    20011004
    EP 1323716
                          A1
                                20030702
                                            EP 2001-972687
                                                                    20011004
             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
     US 2004034049
                                            US 2003-398278
                                                                    20030401
                          A1
                                20040219
PRIORITY APPLN. INFO.:
                                                                 A 20001005
                                             JP 2000-306801
                                            WO 2001-JP8739
                                                                 W 20011004
OTHER SOURCE(S):
                         MARPAT 136:309840
GΙ
```

N R3

AB The title compds. I [R1 and R2 are each H, a hydrocarbon group, a heterocyclic group, or R1 and R2 together with the carbon atom adjacent thereto may form a ring; R3 is H, a hydrocarbon group, or a heterocyclic group; W is R4R5N, etc.; R4 is acyl which is substituted with an arom. group and addnl. bears an optionally substituted aliph. hydrocarbon group or an arom. group; R5 is H, C1-6 alkyl, or acyl; Y is O, S, or NH; and ring C is an optionally substituted benzene ring] are prepd. Three compds. of this invention at 1 μM gave 344% to 478% promotion of neuron generation. Formulations are given.

IT <u>216989-15-0P</u> <u>216989-16-1</u>P <u>216989-18-3</u>P

216989-19-4P 216989-20-7P 216989-21-8P 216989-22-9P 216989-23-0P 216989-24-1P 216989-25-2P 216989-28-5P 216989-29-6P 216989-30-9P 216989-38-7P 216989-39-8P 216989-43-4P 216989-44-5P 216989-46-7P

409366-59-2P 409366-61-6P
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of heterocyclic compds. as promoters for proliferation and differentiation of stem cells and neuron precursor cells)

RN 216989-15-0 HCAPLUS

CN Benzofuran, 2,3-dihydro-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]-5-(phenylmethoxy)- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} & \text{Me} & \text{Me} \\ & \text{Me} & \text{Me} \\ & \text{Ph-CH} \, 2-0 \\ & \text{Me} \end{array}$$

RN 216989-16-1 HCAPLUS

CN Benzenamine, 4-[2,3-dihydro-2,2,4,6,7-pentamethyl-5-(phenylmethoxy)-3-benzofuranyl]-N,N-dimethyl- (9CI) (CA INDEX NAME)

RN <u>216989-18-3</u> HCAPLUS

CN Benzofuran, 2,3-dihydro-5-[(4-methoxyphenyl)methoxy]-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]- (9CI) (CA INDEX NAME)

RN <u>216989-19-4</u> HCAPLUS

CN Benzofuran, 2,3-dihydro-5-[(4-methoxyphenyl)methoxy]-2,2-dimethyl-3-[4-(1-methylethyl)phenyl]- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} \text{Me} & \text{Me} \\ \hline \\ \text{CH}_2 - 0 \\ \hline \end{array}$$

RN 216989-20-7 HCAPLUS

CN Benzenamine, 4-[2,3-dihydro-5-[(4-methoxyphenyl)methoxy]-2,2,4,6,7-pentamethyl-3-benzofuranyl]-N,N-dimethyl- (9CI) (CA INDEX NAME)

RN <u>216989-21-8</u> HCAPLUS

CN Morpholine, 4-[4-[2,3-dihydro-5-[(4-methoxyphenyl)methoxy]-2,2,4,6,7-pentamethyl-3-benzofuranyl]phenyl]- (9CI) (CA INDEX NAME)

RN <u>216989-22-9</u> HCAPLUS

CN Piperazine, 1-[4-[2,3-dihydro-5-[(4-methoxyphenyl)methoxy]-2,2,4,6,7-pentamethyl-3-benzofuranyl]phenyl]-4-methyl- (9CI) (CA INDEX NAME)

RN 216989-23-0 HCAPLUS

CN Benzofuran, 2,3-dihydro-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]-5-[[4-(methylthio)phenyl]methoxy]- (9CI) (CA INDEX NAME)

RN 216989-24-1 HCAPLUS

CN Benzofuran, 2,3-dihydro-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]-5-[[4-(methylsulfinyl)phenyl]methoxy]- (9CI) (CA INDEX NAME)

RN 216989-25-2 HCAPLUS

h

CN Benzofuran, 2,3-dihydro-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]-5-[[4-(methylsulfonyl)phenyl]methoxy]- (9CI) (CA INDEX NAME)

$$\begin{array}{c} \text{Me} - S \\ 0 \\ \text{CH}_2 - 0 \\ \text{Me} \end{array} \begin{array}{c} \text{Me} \\ \text{Me} \\ \text{Me} \end{array}$$

RN 216989-28-5 HCAPLUS

CN Benzofuran, 5-(3,3-diphenylpropoxy)-2,3-dihydro-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]- (9CI) (CA INDEX NAME)

RN 216989-29-6 HCAPLUS

CN Benzoic acid, 4-[[[2,3-dihydro-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]-5-benzofuranyl]oxy]methyl]-, methyl ester (9CI) (CFINDEX NAME)

RN <u>216989-30-9</u> HCAPLUS

CN Benzeneacetic acid, α -[[2,3-dihydro-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]-5-benzofuranyl]oxy]-, methyl ester (9CI) (CA INDEX NAME)

RN 216989-38-7 HCAPLUS

CN Benzofuran, 2,3-dihydro-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]-5-(3-phenylpropoxy)- (9CI) (CA INDEX NAME)

$$\begin{array}{c} \text{Me} \\ \text{Me} \\ \text{Ph-(CH 2)}_{3-0} \\ \text{Me} \end{array}$$

RN 216989-39-8 HCAPLUS

CN Benzofuran, 2,3-dihydro-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]-5-(2-phenylethoxy)- (9CI) (CA INDEX NAME)

RN <u>216989-43-4</u> HCAPLUS

CN Benzofuran, 2,3-dihydro-6-[(4-methoxyphenyl)methoxy]-2,2-dimethyl-3-[4-(1-methylethyl)phenyl]- (9CI) (CA INDEX NAME)

RN 216989-44-5 HCAPLUS

CN Spiro[benzofuran-2(3H), 4'-piperidine], 5-[(4-methoxyphenyl)methoxy]-4,6,7-trimethyl-3-[4-(1-methylethyl)phenyl]-1'-(phenylmethyl)- (9CI) (CA INDEX NAME)

RN 216989-46-7 HCAPLUS

CN Spiro[benzofuran-2(3H),4'-piperidine], 5-[(4-methoxyphenyl)methoxy]1',4,6,7-tetramethyl-3-[4-(1-methylethyl)phenyl]- (9CI) (CA INDEX NAME)

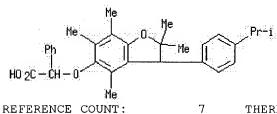
RN 409366-59-2 HCAPLUS

CN Benzofuran, 2,3-dihydro-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]-5-[[(2E)-3-phenyl-2-propenyl]oxy]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

409366-61-6 HCAPLUS RN

Benzeneacetic acid, α -[[2,3-dihydro-2,2,4,6,7-pentamethyl-3-[4-(1-CN methylethyl)phenyl]-5-benzofuranyl]oxy]- (9CI) (CA INDEX NAME)



REFERENCE COUNT:

THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4ANSWER 3 OF 4 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1998:806634 HCAPLUS

DOCUMENT NUMBER: 130:38285

TITLE: Benzofuran derivatives useful for suppressing

neurodegeneration.

INVENTOR(S): Ohkawa, Shigenori; Setoh, Masaki; Kakihana, Mitsuru;

Okura, Masahiro

PATENT ASSIGNEE(S): Takeda Chemical Industries, Ltd., Japan

SOURCE: PCT Int. Appl., 132 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9855454	A2	19981210	WO 1998-JP2482	19980604

h eb c g cg b cg .

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WO 9855454
                           A3
                                 19990304
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             HU, ID, IL, IS, KG, KR, KZ, LC, LK, LR, LT, LV, MD, MG, MK, MN,
             MX, NO, NZ, PL, RO, RU, SG, SI, SK, SL, TJ, TM, TR, TT, UA, US,
             UZ, VN, YU, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
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             CM, GA, GN, ML, MR, NE, SN, TD, TG
     AU 9875503
                           A1
                                 19981221
                                             AU 1998-75503
                                                                      19980604
     JP 1104<u>9765</u>
                           A2
                                 19990223
                                             JP 1998-155709
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                                 20000329
     EP 988289
                          A2
                                             EP 1998-923128
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             IE, FI
PRIORITY APPLN. INFO.:
                                             JP 1997-148325
                                                                  A
                                                                     19970605
                                             WO 1998-JP2482
                                                                     19980604
OTHER SOURCE(S):
                         MARPAT 130:38285
GI
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$$\begin{array}{c} \text{MeO} \\ \text{R3} \\ \text{R4-X} \\ \text{R1} \end{array} \begin{array}{c} \text{Me} \\ \text{Me} \\ \text{Me} \end{array} \begin{array}{c} \text{Me} \\ \text{Me} \\ \text{Me} \end{array} \begin{array}{c} \text{Me} \\ \text{Me} \\ \text{Me} \end{array}$$

Title compds. I [R1, R2 = H, (un)substituted hydrocarbon group; or R1 and R2 form a 3- to 8-membered carbo- or heterocyclic ring which may be substituted; R3 = H, (un)substituted lower alkyl or arom. group; R4 = (un)substituted arom. or araliph. group, or acyl; X , Y = O or S which may be oxidized; benzene ring may be further substituted] and their salts are disclosed. The compds. suppress β -amyloid toxicity, and are thus useful as agents for treating of preventing neurodegenerative diseases such as Alzheimer's disease or Parkinsonism. Prepns. of 33 compds. I and their intermediates are described. For instance, etherification of 3-(4-isopropylphenyl)-2,2,4,6,7-pentamethyl-2,3-dihydrobenzofuran-5-ol with 4-methoxybenzyl chloride using NaH in DMF gave 49% title compd. II. Seven example compds. gave 27.3-47.0% in vitro protection of human neuroblastoma SK-N-SH cells from β -amyloid neurotoxicity.

neurodegeneration)

RN 216989-23-0 HCAPLUS

CN Benzofuran, 2,3-dihydro-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]-5-[[4-(methylthio)phenyl]methoxy]- (9CI) (CA INDEX NAME)

RN <u>216989-24-1</u> HCAPLUS

CN Benzofuran, 2,3-dihydro-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]-5-[[4-(methylsulfinyl)phenyl]methoxy]- (9CI) (CA INDEX NAME)

RN 216989-26-3 HCAPLUS

CN Benzofuran, 2,3-dihydro-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]-5-[(3-phenyl-2-propenyl)oxy]- (9CI) (CA INDEX NAME)

$$\begin{array}{c} \text{Me} \\ \text{Me} \\ \text{Ph-CH} = \text{CH-CH}_2 - 0 \\ \text{Me} \end{array}$$

RN 216989-30-9 HCAPLUS

CN Benzeneacetic acid, α -[[2,3-dihydro-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]-5-benzofuranyl]oxy]-, methyl ester (9CI) (CA INDEX NAME)

IT 216989-15-0P, 5-(Benzyloxy)-3-(4-isopropylphenyl)-2,2,4,6,7pentamethyl-2,3-dihydrobenzofuran 216989-16-1P,
5-(Benzyloxy)-3-[4-(dimethylamino)phenyl]-2,2,4,6,7-pentamethyl-2,3dihydrobenzofuran 216989-18-3P, 3-(4-Isopropylphenyl)-5-[(4-

```
methoxybenzyl)oxy]-2,2,4,6,7-pentamethyl-2,3-dihydrobenzofuran
216989-19-4P, 3-(4-Isopropylphenyl)-5-[(4-methoxybenzyl)oxy]-2,2-
dimethyl-2, 3-dihydrobenzofuran 216989-20-7P,
3-[4-(Dimethylamino)phenyl]-5-[(4-methoxybenzyl)oxy]-2,2,4,6,7-pentamethyl-
2,3-dihydrobenzofuran 216989-21-8P, 5-[(4-Methoxybenzyl)oxy]-3-
[4-(4-morpholinyl)phenyl]-2,2,4,6,7-pentamethyl-2,3-dihydrobenzofuran
216989-22-9P, 5-[(4-Methoxybenzyl)oxy]-2, 2, 4, 6, 7-pentamethyl-3-[4-
(4-methyl-1-piperazinyl)phenyl]-2,3-dihydrobenzofuran 216989-25-2P
, 3-(4-Isopropylphenyl)-2,2,4,6,7-pentamethyl-5-[[4-
(methylsulfonyl)benzyl]oxy]-2,3-dihydrobenzofuran 216989-28-5P,
5-[(3,3-Diphenylpropyl)oxy]-3-(4-isopropylphenyl)-2,2,4,6,7-pentamethyl-
2,3-dihydrobenzofuran 216989-29-6P, Methyl 4-[[[3-(4-
isopropylphenyl)-2,2,4,6,7-pentamethyl-2,3-dihydrobenzofuran-5-
yl]oxy]methyl]benzoate 216989-36-5P 216989-37-6P
216989-38-7P, 3-(4-Isopropylphenyl)-2,2,4,6,7-pentamethyl-5-[(3-
phenylpropyl)oxy]-2,3-dihydrobenzofuran 216989-39-8P,
3-(4-Isopropylphenyl)-2,2,4,6,7-pentamethyl-5-[(2-phenylethyl)oxy]-2,3-
dihydrobenzofuran 216989-43-4P, 3-(4-Isopropylphenyl)-6-[(4-
methoxybenzyl)oxy]-2,2-dimethyl-2,3-dihydrobenzofuran 216989-44-5p
, 1'-Benzyl-3-(4-isopropylphenyl)-5-[(4-methoxybenzyl)oxy]-4,6,7-
trimethylspiro[benzofuran-2(3H),4'-piperidine] 216989-46-7P,
3-(4-Isopropylphenyl)-5-[(4-methoxybenzyl)oxy]-1', 4, 6, 7-
tetramethylspiro[benzofuran-2(3H),4'-piperidine]
RL: BAC (Biological activity or effector, except adverse); BSU (Biological
study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use);
BIOL (Biological study); PREP (Preparation); USES (Uses)
   (product; prepn. of benzofuran derivs. as agents for suppressing
   neurodegeneration)
216989-15-0 HCAPLUS
Benzofuran, 2,3-dihydro-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]-
5-(phenylmethoxy)- (9CI) (CA INDEX NAME)
```

$$\begin{array}{c} \text{Me} \\ \text{Me} \\ \text{Ph-CH}_{\,2}-0 \\ \text{Me} \end{array} \begin{array}{c} \text{Me} \\ \text{Me} \\ \text{Pr-i} \\ \text{Me} \end{array}$$

RN <u>216989-16-1</u> HCAPLUS

RN

CN

h

CN Benzenamine, 4-[2,3-dihydro-2,2,4,6,7-pentamethyl-5-(phenylmethoxy)-3-benzofuranyl]-N,N-dimethyl- (9CI) (CA INDEX NAME)

RN 216989-18-3 HCAPLUS

CN Benzofuran, 2,3-dihydro-5-[(4-methoxyphenyl)methoxy]-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]- (9CI) (CA INDEX NAME)

RN <u>216989-19-4</u> HCAPLUS

CN Benzofuran, 2,3-dihydro-5-[(4-methoxyphenyl)methoxy]-2,2-dimethyl-3-[4-(1-methylethyl)phenyl]- (9CI) (CA INDEX NAME)

RN <u>216989-20-7</u> HCAPLUS

CN Benzenamine, 4-[2,3-dihydro-5-[(4-methoxyphenyl)methoxy]-2,2,4,6,7-pentamethyl-3-benzofuranyl]-N,N-dimethyl- (9CI) (CA INDEX NAME)

RN <u>216989-21-8</u> HCAPLUS

CN Morpholine, 4-[4-[2,3-dihydro-5-[(4-methoxyphenyl)methoxy]-2,2,4,6,7-pentamethyl-3-benzofuranyl]phenyl]- (9CI) (CA INDEX NAME)

RN 216989-22-9 HCAPLUS

CN Piperazine, 1-[4-[2,3-dihydro-5-[(4-methoxyphenyl)methoxy]-2,2,4,6,7-pentamethyl-3-benzofuranyl]phenyl]-4-methyl- (9CI) (CA INDEX NAME)

RN <u>216989-25-2</u> HCAPLUS

CN Benzofuran, 2,3-dihydro-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]-5-[[4-(methylsulfonyl)phenyl]methoxy]- (9CI) (CA INDEX NAME)

RN 216989-28-5 HCAPLUS

CN Benzofuran, 5-(3,3-diphenylpropoxy)-2,3-dihydro-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]- (9CI) (CA INDEX NAME)

$$\begin{array}{c} \text{Me} \\ \text{Me} \\ \text{Ph}_2\text{CH} - \text{CH}_2 - \text{CH}_2 - \text{O} \\ \text{Me} \\ \end{array}$$

RN <u>216989-29-6</u> HCAPLUS

CN Benzoic acid, 4-[[[2,3-dihydro-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]-5-benzofuranyl]oxy]methyl]-, methyl ester (9CI) (CA INDEX NAME)

$$\begin{array}{c} \text{Me} \\ \text{Me} \\$$

RN 216989-36-5 HCAPLUS

CN Benzeneacetic acid, $\alpha-[[(3R)-2,3-dihydro-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]-5-benzofuranyl]oxy]-, (<math>\alpha$ R)-rel-(9CI) (CA INDEX NAME)

eb

Relative stereochemistry.

RN <u>216989-37-6</u> HCAPLUS

CN Benzeneacetic acid, α -[[(3R)-2,3-dihydro-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]-5-benzofuranyl]oxy]-, (α S)-rel-(9CI) (CA INDEX NAME)

Relative stereochemistry.

RN 216989-38-7 HCAPLUS

CN Benzofuran, 2,3-dihydro-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]-5-(3-phenylpropoxy)- (9CI) (CA INDEX NAME)

RN <u>216989-39-8</u> HCAPLUS

CN Benzofuran, 2,3-dihydro-2,2,4,6,7-pentamethyl-3-[4-(1-methylethyl)phenyl]-5-(2-phenylethoxy)- (9CI) (CA INDEX NAME)

$$\begin{array}{c} \text{Me} \\ \text{Me} \\ \text{Ph-CH}_2\text{-CH}_2\text{-0} \\ \text{Me} \end{array}$$

RN 216989-43-4 HCAPLUS

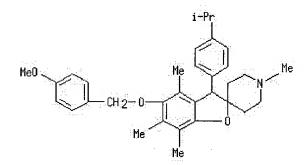
CN Benzofuran, 2,3-dihydro-6-[(4-methoxyphenyl)methoxy]-2,2-dimethyl-3-[4-(1-methylethyl)phenyl]- (9CI) (CA INDEX NAME)

RN <u>216989-44-5</u> HCAPLUS

CN Spiro[benzofuran-2(3H),4'-piperidine], 5-[(4-methoxyphenyl)methoxy]-4,6,7-trimethyl-3-[4-(1-methylethyl)phenyl]-1'-(phenylmethyl)- (9CI) (CA INDEX NAME)

RN <u>216989-46-7</u> HCAPLUS

CN Spiro[benzofuran-2(3H),4'-piperidine], 5-[(4-methoxyphenyl)methoxy]1',4,6,7-tetramethyl-3-[4-(1-methylethyl)phenyl]- (9CI) (CA INDEX NAME)



L4 ANSWER 4 OF 4 HCAPLUS COPYRIGHT 2004 ACS on STN

Full sister Text Sections

ACCESSION NUMBER: 1988:549335 HCAPLUS

DOCUMENT NUMBER: 109:149335

TITLE: Preparation of 5-hydroxycoumaran derivatives as

cardiovascular and antiallergy agents

INVENTOR(S): Terao, Shinji; Maki, Yoshitaka

PATENT ASSIGNEE(S): Takeda Chemical Industries, Ltd., Japan

SOURCE: Eur. Pat. Appl., 39 pp.

CODEN: EPXXDW

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.		KINI	D DATE	APPLICATION NO.		DATE
			·			
EP 273647		A1	19880706	EP 1987-311122		19871217
EP 273647		В1	19920311			
R: AT,	BE, CH,	DE,	ES, FR, GB,	GR, IT, LI, LU, NL, SE	2	
JP 01272578		A2	19891031	JP 1987-310346		19871207
JP 08005871		В4	19960124			
AT 73448		E	19920315	AT 1987-311122		19871217
DK 8706789		A	19880628	DK 1987-6789		19871222
<u>US 4857516</u>		A	19890815	US 1987-136273		19871222
HU 48609		A2	19890628	HU 1987-5988		19871223
<u>HU 206332</u>		В	19921028			
AU 8783040		A1	19880630	AU 1987-83040		19871224
AU 605818		В2	19910124			
CA 1325635		A1	19931228	CA 1987-555354		19871224
PRIORITY APPLN. I	NFO.:			JP 1986-313380	Α	19861227
				JP 1987-235491	A	19870918

EP 1987-311122

A 19871217

eb

OTHER SOURCE(S):

CASREACT 109:149335; MARPAT 109:149335

$$R^{00}$$
 R^{1}
 R^{2}
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 $R^{$

The title compds. [I; R = alkyl; R0 = H, acyl; R1-R4 = (un)substituted alkyl; R1R2 = CH:CHCH:CH; R3R4 = polymethylene; R5 = (un)substituted alkyl, aryl, heterocyclyl] were prepd. 4-FC6H4COCHMe2 (prepn. given) was added to 1-bromo-2,5-dimethoxy-3,4,6-trimethylbenzene in THF previously treated with BuLi and the mixt. stirred 1 h to give 92.3% diphenylpropanol II which was refluxed 18 h in 47 wt.% aq. HBr to give 74.8% title compd. III. The latter, at 100 mg/kg orally gave 93% inhibition of the excitatory behavior induced by spinal intrathecal injection of FeC12 soln. in mice.

IT 116674-58-9P

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. of, as cardiovascular and antiallergic agent)

RN <u>116674-58-9</u> HCAPLUS

CN 5-Benzofuranol, 3-(4-fluorophenyl)-2,3-dihydro-2,2,4,6,7-pentamethyl-, benzoate (9CI) (CA INDEX NAME)

=> file caold COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 23.76 181.49 DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION CA SUBSCRIBER PRICE -2.80-2.80

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FILE COVERS 1907-1966

FILE LAST UPDATED: 01 May 1997 (19970501/UP)

This file contains CAS Registry Numbers for easy and accurate substance identification. Title keywords, authors, patent assignees, and patent information, e.g., patent numbers, are now searchable from 1907-1966. TIFF images of CA abstracts printed between 1907-1966 are available in the PAGE display formats.

This file supports REG1stRY for direct browsing and searching of all substance data from the REGISTRY file. Enter $\underline{\text{HELP FIRST}}$ for more information.

=> d his

(FILE 'HOME' ENTERED AT 12:56:20 ON 16 DEC 2004)

FILE 'REGISTRY' ENTERED AT 12:56:26 ON 16 DEC 2004

L1 STRUCTURE UPLOADED

L2 1 S L1

L3 24 S L1 FULL

FILE 'HCAPLUS' ENTERED AT 12:59:53 ON 16 DEC 2004 4 S L3

FILE 'CAOLD' ENTERED AT 13:01:09 ON 16 DEC 2004

=> s 1.3

L4

L5 0 L3

=> file medline, biosis, embase, caplus

COST IN U.S. DOLLARS SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST 0.84 182.33

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE TOTAL

ENTRY SESSION

CA SUBSCRIBER PRICE

0.00 -2.80

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=> s ohkawa, s?au

h

'?' TRUNCATION SYMBOL NOT VALID WITHIN 'S?AU'
The truncation symbol ? may be used only at the end of a search
term. To specify a variable character within a word use '!', e.g.,
'wom!n' to search for both 'woman' and 'women'. Enter "HELP

TRUNCATION" at an arrow prompt (=>) for more information.

=> s ohkawa, s?/au

L6 1169 OHKAWA, S?/AU

=> s setoh, m?/au

20 SETOH, M?/AU

=> s kakihana, m?/au

709 KAKIHANA, M?/AU

=> s okura, m?/au

L9 293 OKURA, M?/AU

=> s 16 and 17 and 18 and 19

1 L6 AND L7 AND L8 AND L9

=> d l10, ibib abs fhitstr, 1

L10 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2004 ACS on STN

References Text

ACCESSION NUMBER:

1998:806634 CAPLUS

DOCUMENT NUMBER:

130:38285

TITLE:

Benzofuran derivatives useful for suppressing

neurodegeneration.

INVENTOR(S):

Ohkawa, Shigenori; Setoh, Masaki; Kakihana,

Mitsuru; Okura, Masahiro

PATENT ASSIGNEE(S):

SOURCE:

Takeda Chemical Industries, Ltd., Japan

PCT Int. Appl., 132 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PA:	rent :	NO.			KIN	D	DATE			APPL	ICAT	ION	NO.		D	ATE	
						-									-		
\overline{MO}	9855	<u>454</u>			A2		1998	1210		<u>wo 1</u>	998-	JP24	<u>82</u>	-	1	9980	604
WO	9855	454			A3		1999	0304									
	w:	AL,	AM,	ΑU,	AZ,	BA,	BB,	BG,	BR,	BY,	CA,	CN,	CU,	CZ,	EE,	GE,	GW,
		ΗU,	ID,	ΙL,	IS,	KG,	KR,	ΚZ,	LC,	LK,	LR,	LT,	LV,	MD,	MG,	MK,	MN,
		MX,	NO,	NZ,	PL,	RO,	RU,	SG,	SI,	SK,	SL,	ТJ,	TM,	TR,	TT,	UA,	US,
		UZ,	VN,	YU,	AM,	AZ,	BY,	KG,	ΚZ,	MD,	RU,	ТJ,	MT				
	RW:	GH,	GM,	ΚE,	LS,	MW,	SD,	SZ,	UG,	ZW,	AT,	BE,	CH,	CY,	DE,	DK,	ES,
		FI,	FR,	GB,	GR,	ΙE,	IT,	LU,	MC,	NL,	PT,	SE,	BF,	ВJ,	CF,	CG,	CI,
		CM,	GΑ,	GN,	ML,	MR,	NE,	SN,	TD,	TG							
AU	9875	503			A1		1998	1221		AU 1	998-	7550	3		1	9980	604
JP	1104	9765			A2		1999	0223		JP 1	998-	1557	09		1	9980	604
EP	9882	89			A2		2000	0329		EP 1	998-	9231.	28		1	9980	604
 -	R:	ΑT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,
		ΙE,	FΙ												2		
PRIORITY	APP	LN.	INFO	. :						JP 1:	997-	1483	25		A 1	9970	605
										WO 1	998-	JP24	82	,	W 1	9980	604
OTHER SO	OURCE	(S):			MAR	PAT	130:	38285	5				-				

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Title compds. I [R1, R2 = H, (un)substituted hydrocarbon group; or R1 and R2 form a 3- to 8-membered carbo- or heterocyclic ring which may be substituted; R3 = H, (un)substituted lower alkyl or arom. group; R4 = (un)substituted arom. or araliph. group, or acyl; X , Y = O or S which may be oxidized; benzene ring may be further substituted] and their salts are disclosed. The compds. suppress β -amyloid toxicity, and are thus useful as agents for treating of preventing neurodegenerative diseases such as Alzheimer's disease or Parkinsonism. Prepns. of 33 compds. I and their intermediates are described. For instance, etherification of 3-(4-isopropylphenyl)-2,2,4,6,7-pentamethyl-2,3-dihydrobenzofuran-5-ol with 4-methoxybenzyl chloride using NaH in DMF gave 49% title compd. II. Seven example compds. gave 27.3-47.0% in vitro protection of human neuroblastoma SK-N-SH cells from β -amyloid neurotoxicity.

=> d his

(FILE 'HOME' ENTERED AT 12:56:20 ON 16 DEC 2004)

FILE 'REGISTRY' ENTERED AT 12:56:26 ON 16 DEC 2004

L1 STRUCTURE UPLOADED

L2 1 S L1

L3 24 S L1 FULL

FILE 'HCAPLUS' ENTERED AT 12:59:53 ON 16 DEC 2004

L4 4 S L3

FILE 'CAOLD' ENTERED AT 13:01:09 ON 16 DEC 2004

L5 0 S L3

FILE 'MEDLINE, BIOSIS, EMBASE, CAPLUS' ENTERED AT 13:02:10 ON 16 DEC 2004

L6 1169 S OHKAWA, S?/AU

L7 20 S SETOH, M?/AU

L8 709 S KAKIHANA, M?/AU

L9 293 S OKURA, M?/AU

L10 1 S L6 AND L7 AND L8 AND L9

=> s 16 and 17

L11 6 L6 AND L7

=> s 111 and 18

L12 1 L11 AND L8

=> d lll, ibib abs fhitstr, 1-6

L11 ANSWER 1 OF 6 BIOSIS COPYRIGHT (c) 2004 The Thomson Corporation. on STN



h ebc gcgb cg

ACCESSION NUMBER: 2003:53330 BIOSIS DOCUMENT NUMBER: PREV200300053330

TITLE: Benzofuran derivatives, process for the preparation of the

same and uses thereof.

AUTHOR(S): Ohkawa, Shigenori [Inventor, Reprint Author]; Arikawa,

Yasuyoshi [Inventor]; Kato, Kouki [Inventor]; Okura,

Masahiro [Inventor]; Setoh, Masaki [Inventor]

CORPORATE SOURCE: Takatsuki, Japan

ASSIGNEE: Takeda Chemical Industries, Ltd., Osaka, Japan

PATENT INFORMATION: US 6479536 November 12, 2002

SOURCE: Official Gazette of the United States Patent and Trademark

Office Patents, (Nov 12 2002) Vol. 1264, No. 2. http://www.uspto.gov/web/menu/patdata.html. e-file.

ISSN: 0098-1133 (ISSN print).

DOCUMENT TYPE: Patent LANGUAGE: English

ENTRY DATE: Entered STN: 22 Jan 2003

Last Updated on STN: 22 Jan 2003

Compounds represented by the formula: ##STR1## wherein R1 and R2 are AΒ hydrogen atom, a hydrocarbon group or a heterocyclic group, or R1 and R2 may form, together with the adjacent carbon atom, a 3- to 8-membered homocyclic or heterocyclic ring, W indicates (i) a group represented by the formula: ##STR2## wherein ring B indicates a 5- to 7-membered ring, or (ii) a group represented by the formula: ##STR3## wherein R4 indicates (1) an aliphatic hydrocarbon group, which may be substituted with an aromatic group, or (2) an acyl group containing an aromatic group, R5 is hydrogen atom, a C1-6 alkyl, or an acyl group, provided that, when W is Wa, R3 is hydrogen atom, a hydrocarbon group or a heterocyclic group, when W is Wb, R3 indicates a C6-14 aryl group, or salts thereof or prodrugs thereof have an excellent action to inhibit neurodegeneration and the like as well as an excellent brain penetrability and are low in the toxicity, thereby being useful as prophylactic or therapeutic drugs for nerve degenerative diseases and the like.

L11 ANSWER 2 OF 6 BIOSIS COPYRIGHT (c) 2004 The Thomson Corporation. on STN

FUI CERSION NUMBER :

ACCESSION NUMBER: 2002:447093 BIOSIS DOCUMENT NUMBER: PREV200200447093

TITLE: Tricyclic compound, their production and use.

AUTHOR(S): Ohkawa, Shigenori [Inventor, Reprint author]; Setoh,

Masaki [Inventor]; Terashita, Zen-ichi [Inventor]

CORPORATE SOURCE: Takatsuki, Japan

ASSIGNEE: Takeda Chemical Industries, Ltd., Osaka, Japan

PATENT INFORMATION: US 6417213 July 09, 2002

SOURCE: Official Gazette of the United States Patent and Trademark

Office Patents, (July 9, 2002) Vol. 1260, No. 2. http://www.uspto.gov/web/menu/patdata.html. e-file.

CODEN: OGUPE7. ISSN: 0098-1133.

DOCUMENT TYPE: Patent LANGUAGE: English

ENTRY DATE: Entered STN: 21 Aug 2002

Last Updated on STN: 21 Aug 2002

AB A compound of the formula ##STR1## wherein R1 is H or a substituent; m is 1-3; Ar is an aromatic group which may be substituted; X is a bond or a divalent straight-chain group having 1-6 atoms which may be substituted; Y is --S--, --O--, or --N(R2)-- (R2 is H or a substituent group), Z is --Ndbd or --C(R3)dbd (R3 is H or a hydrocarbon group), ring A is a benzene ring; ring B is a 5- to 7-membered ring which may be substituted, or a salt thereof is useful for eliciting a prostaglandin I2 receptor agonistic

effect.

ANSWER 3 OF 6 BIOSIS COPYRIGHT (c) 2004 The Thomson Corporation. on STN

Text Reterences

2001:416973 BIOSIS ACCESSION NUMBER: DOCUMENT NUMBER: PREV200100416973

TITLE:

Tricyclic compounds, their production and use.

AUTHOR (S):

Ohkawa, Shigenori [Inventor, Reprint author]; Setoh,

Masaki [Inventor]; Terashita, Zen-ichi [Inventor]

CORPORATE SOURCE:

Takatsuki, Japan

ASSIGNEE: Takeda Chemical Industries, Ltd., Osaka, Japan

PATENT INFORMATION: US 6248766 June 19, 2001

SOURCE:

Official Gazette of the United States Patent and Trademark Office Patents, (June 19, 2001) Vol. 1247, No. 3. e-file.

CODEN: OGUPE7. ISSN: 0098-1133.

DOCUMENT TYPE:

Patent English

LANGUAGE: ENTRY DATE:

Entered STN: 29 Aug 2001

Last Updated on STN: 22 Feb 2002

AB A compound of the formula: ##STR1## wherein R1 is H or a substituent; m is 1-3; Ar is an aromatic group which may be substituted; X is a bond or a divalent straight-chain group having 1-6 atoms which may be substituted; Y is --S--, --O--, or --N(R2 -- (R2 is H or a substituent group), Z is --Ndbd or --C(R3)dbd (R3 is H or a hydrocarbon group), ring A is a benzene ring; ring B is a 5- to 7-membered ring which may be substituted, or a salt thereof is useful for eliciting a prostaglandin I2 receptor agonistic effect.

L11 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN

Text

ACCESSION NUMBER: 2000:401810

DOCUMENT NUMBER:

TITLE:

INVENTOR(S):

CAPLUS

133:43436

and preventives for neurodegeneration

Ohkawa, Shigenori; Arikawa, Yasuyoshi; Kato, Kouki;

Preparation of benzofuran derivatives as inhibitors

Okura, Masahiro; Setoh, Masaki

PATENT ASSIGNEE(S):

Takeda Chemical Industries, Ltd., Japan

SOURCE:

PCT Int. Appl., 247 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

Japanese

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PA	rent :	NO.			KIN	D	DATE			APPL	ICAT	ION	NO.		D	ATE	
						_									-		
MO	2000	0342	<u>62</u>		A1		2000	0615		WO 1	999-	JP67	<u>64</u>		1	9991	202
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		SL,	ТJ,	TM,	TR,	TT,	TZ,	UA,	US,	UZ,	VN,	YU,	ZA,	AM,	AZ,	BY,	KG,
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EP	1136		A1		2001	0926		EP 1	999-	9732	89		1	9991:	202		
EΡ	1136	477			В1		2004	0310					_				

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	IE,	SI,	LT,	LV,	FI,	RO										
<u>AT 261</u>	<u>435</u>			\mathbf{E}		2004	0315	Ī	TA	1999	9732	89			19991	202
ES 221	3407			тЗ		2004	0816]	ES :	1999-	9732	89			19991	202
PT 113	6477			\mathbf{T}		2004	0831]	PT :	1999-	9732	<u>89</u>			19991	202
JP 200	02263	88		A2		2000	0815	į	JP :	1.999-	3443	45			19991	.203
JP 355	3442			В2		2004	0811									•
JP 200	21610	88		A2		2002	0604	3	JP 2	2001-	3140	<u> 27</u>			19991	.203
NO 200	10027	<u> 26</u>		A		2001	0731	Ţ	NO 2	2001-	2726				20010	601
<u>US 647</u>	9536			В1		2002	1112	Ţ	US 2	2001-	8572	93			20010	601
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						•		Ī	WO :	1999-	JP67	64	I	W	19991	202
								<u>.</u>	JP :	1999-	3443	<u>45</u>	7	EA	19991	.203
								Ţ	US 2	2001-	8572	93	1	A3	20010	601
	_ , _ ,							_								

OTHER SOURCE(S):

MARPAT 133:43436

GT For diagram(s), see printed CA Issue.

AΒ Compds. represented by general formula (I), salts of the same, or prodrugs of both [wherein R1 and R2 are each hydrogen, hydrocarbyl or a heterocyclic group, or alternatively R1 and R2 together with the carbon atom adjacent thereto may form a three- to eight-membered homo- or heterocyclic ring; W is (i) a group represented by general formula Q: [wherein B is a five- to seven-membered ring], or (ii) a group represented by general formula R4R5N [wherein R4 is (1) aliph. hydrocarbyl substituted with an arom. group or (2) acyl bearing an arom. group; and R5 is hydrogen, C1-6 alkyl or acyl]; and when W is Q, R3 is hydrogen, hydrocarbyl or a heterocyclic group, whereas when W is R4R5N, R3 is C6-14 aryl] are prepd. These compds. exhibit excellent effects of inhibiting nerve degeneration and toxicity of β -amyloid and excellent activity like nerve nutritional factor and possess intracerebral transmigration properties, and low toxicity, thus being useful as preventive and therapeutic agents for nerve degeneration diseases such as Alzheimer's disease and Parkinson's disease. Thus, a mixt. of 2,2,4,6,7-pentamethyl-3-(4-methylphenyl)-2,3-dihydro-1-benzofuran-5-amine (prepn. given), 1,2-bis(chloromethyl)-4,5-dimethoxybenzene, Na2CO3, and tetrabutylammonium iodide in THF was refluxed for 11 h to give 16% 5,6-dimethoxy-2-[2,2,4,6,7pentamethyl-3-(4-methylphenyl)-2,3-dihydro-1-benzofuran-5-yl]isoindoline (II). II in vitro showed 28.2% cytoprotective activity against LY-294002-induced cytotoxicity in SK-N-SH cells. Pharmaceutical formulations contg. I were also prepd.

REFERENCE COUNT:

9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 5 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN L11

FUI Text Pererences ACCESSION NUMBER:

1998:806634 CAPLUS

DOCUMENT NUMBER: 130:38285

TITLE: Benzofuran derivatives useful for suppressing

neurodegeneration.

INVENTOR (S): Ohkawa, Shigenori; Setoh, Masaki; Kakihana,

Mitsuru; Okura, Masahiro

PATENT ASSIGNEE(S): Takeda Chemical Industries, Ltd., Japan

PCT Int. Appl., 132 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

SOURCE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT	no.			KIN	D	DATE			APPL	ICAT	ION :	NO.		D	ATE	
WO 985				A2	_	 1998	 1210	a	wo 1	998-	JP24	8 <u>2</u>		1	9980	604
<u>wo 985</u>	55454			АЗ		1999	0304									
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JP 110	049765			A2		1999	0223		JP 1	998-	1557	_ 09		1	9980	604
EP 988	3289	•		A2		2000	0329	•	EP 1	998-	9231	28		1	9980	604
R	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,
	IE,	FI														
PRIORITY A	PPLN.	INFO	.:						JP 1	997	1483	25		A 1	9970	605
									wo 1	998-	JP24	82	,	W 1	9980	604
OTHER SOURG	CE(S):			MAR	PAT	130:	3828	5								

Me0 R3 11

Title compds. I [R1, R2 = H, (un)substituted hydrocarbon group; or R1 and AΒ R2 form a 3- to 8-membered carbo- or heterocyclic ring which may be substituted; R3 = H, (un)substituted lower alkyl or arom. group; R4 = (un) substituted arom. or araliph. group, or acyl; X, Y = O or S which may be oxidized; benzene ring may be further substituted] and their salts are disclosed. The compds. suppress β -amyloid toxicity, and are thus useful as agents for treating of preventing neurodegenerative diseases such as Alzheimer's disease or Parkinsonism. Prepns. of 33 compds. I and their intermediates are described. For instance, etherification of 3-(4-isopropylphenyl)-2,2,4,6,7-pentamethyl-2,3-dihydrobenzofuran-5-ol with 4-methoxybenzyl chloride using NaH in DMF gave 49% title compd. II. Seven example compds. gave 27.3-47.0% in vitro protection of human neuroblastoma SK-N-SH cells from β -amyloid neurotoxicity.

L11 ANSWER 6 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN



ACCESSION NUMBER:

DOCUMENT NUMBER:

TITLE:

1998:208535 CAPLUS

128:257432

Preparation of tricyclic compounds as prostaglandin I2

receptor agonists

Ohkawa, Shigenori; Setoh, Masaki; Terashita, Zen-ichi

Takeda Chemical Industries, Ltd., Japan

PCT Int. Appl., 151 pp.

CODEN: PIXXD2

INVENTOR(S):

PATENT ASSIGNEE(S):

SOURCE:

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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\mathbf{T}	W 4169	53			В		2001	0101		TW 1	997-	8611	3705			19	970	920
<u>C2</u>	A 2264	641			AA		1998	0402		CA 1	997-	2264	641			19	970	924
A	J 9743	<u>973</u>			A1		1998	0417		AU 1	997-	4397	3			19	970	924
J	P 1015 P 9295	2480			A2		1998	0609	,	JP 1	997-	2574	80			19	970	924
\mathbf{E}	P 9295	<u>34</u>			A1		1999	0721	`	EP 1	997-	9421	96			19	970	924
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U:	5 2002	0069	44		A1		2002	0117		US 2	001-	8009	88			20	010	307
U	5 6417	213			B2		2002	0709										
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										WO 1	997-	JP33	84		W	19	970	924
										US 1	999-	2544	<u>46</u>		A3	19	990	309
OTHER S	SOURCE	(S):			MAR	TAG	128:	2574	32									
GI								-	- 00-									000

$$\begin{array}{c} Z = X - Ar \\ Z = Y \\ R^{1}C - |CH| 2 - 0 - A - B \end{array}$$

AB The title compds. [I; Rl = H, a substituent; m = 1-3; Ar = (un) substituted arom. group; X = a bond, (un) substituted divalent straight-chain group having 1-6 atoms; Y = S, O, N(R2) (R2 = H, a substituent); Z = N, C(R3) (R3 = H, a hydrocarbon); ring A = a benzene ring; ring B = (un) substituted 5-7 membered ring], useful for eliciting a prostaglandin I2 receptor agonistic effect, inhibiting a platelet aggregation, and for the

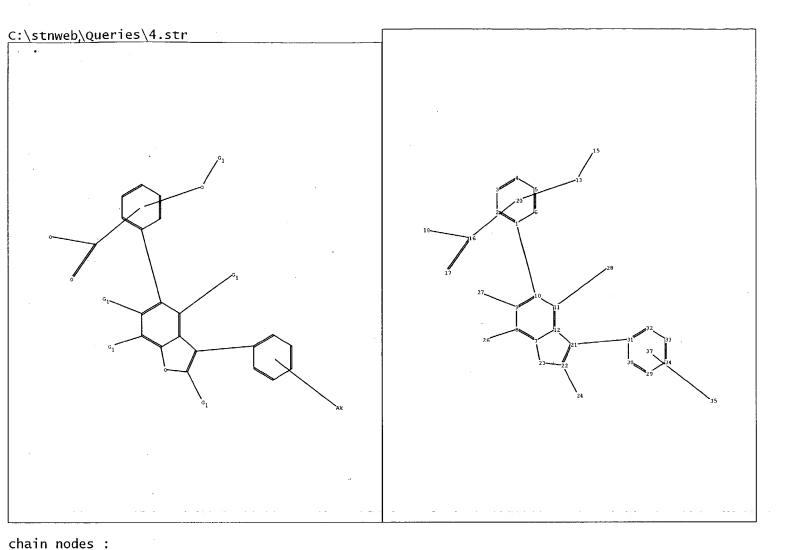
prophylaxis or treatment of transient ischemic attack, diabetic neuropathy, peripheral vascular diseases or ulcer, were prepd. and formulated. Thus, reaction of Et [(2-mercapto-4,5-dihydronaphtho[1,2-d]thiazol-6-yl)oxy]acetate with 2,2-diphenylethyl methanesulfonate in the presence of K2CO3 in DMF followed by hydrolysis the resulting Et {[2-(2,2-diphenylethyl)thio-4,5-dihydronaphtho[1,2-d]thiazol-6-yl]oxy}acetate with 1N NaOH afforded 61% II which showed IC50 of 0.024 μM against PGI2 receptor binding, and IC50 of 0.54 μM against platelet aggregation.

13

REFERENCE COUNT:

THERE ARE 13 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

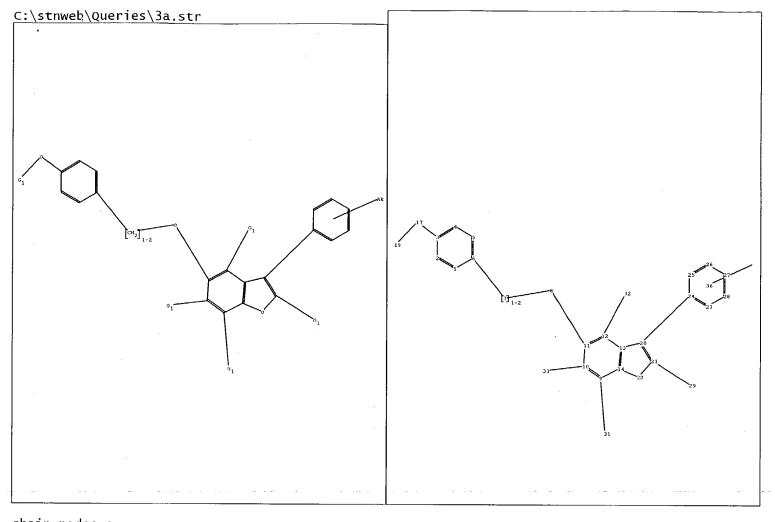
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```
13 15 16 17 18 24 26 27 28 35
ring nodes :
   1 2 3 4 5 6 7 8
                          9 10 11 12 21 22 23 29 30 31 32 33 34
chain bonds :
   1-10 8-26 9-27 11-28 13-15 16-18 16-17 21-31 22-24
ring bonds :
   1-2 1-6 2-3 3-4 4-5 5-6 7-8 7-12 7-23 8-9 9-10 10-11 11-12 12-21 21-22 22-23 29-30 29-34 30-31 31-32 32-33 33-34
exact/norm bonds :
    8-26 9-27 11-28 13-15 16-18 16-17 22-24
exact bonds:
   1-10 7-23 12-21 21-22 21-31 22-23
normalized bonds:
1-2 1-6 2-3 3-4 4-5 5-6 7-8 7-12 8-9 9-10 10-11 11-12 29-30 29-34 30-31
    31-32 32-33 33-34
isolated ring systems :
   containing 1 : 7 : 29 :
```

G1:CH3,Et

Match level:
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom 11:Atom 12:Atom 13:CLASS 15:CLASS 16:CLASS 17:CLASS 18:CLASS 19:CLASS 20:CLASS 21:Atom 22:Atom 23:Atom 24:CLASS 26:CLASS 27:CLASS 28:CLASS 29:Atom 30:Atom 31:Atom 32:Atom 33:Atom 34:Atom 35:CLASS 37:CLASS



chain nodes :
 7 8 17 19 29 31 32 33 35
ring nodes :
 1 2 3 4 5 6 9 10 11 12 13 14 20 21 22 23 24 25 26 27 28
chain bonds :
 3-17 6-7 7-8 8-11 9-31 10-33 12-32 17-19 20-24 21-29
ring bonds :
 1-2 1-6 2-3 3-4 4-5 5-6 9-10 9-14 10-11 11-12 12-13 13-14 13-20 14-22 20-21 21-22 23-24 23-28 24-25 25-26 26-27 27-28
exact/norm bonds :
 3-17 8-11 9-31 10-33 12-32 17-19 21-29
exact bonds :
 6-7 7-8 13-20 14-22 20-21 20-24 21-22
normalized bonds :
 1-2 1-6 2-3 3-4 4-5 5-6 9-10 9-14 10-11 11-12 12-13 13-14 23-24 23-28 24-25 25-26 26-27 27-28
isolated ring systems :

G1:CH3,Et

containing 1 : 9 : 23 :

Match level:
 1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:CLASS 8:CLASS 9:Atom 10:Atom 11:Atom 12:Atom 13:Atom 14:Atom 17:CLASS 19:CLASS 20:Atom 21:Atom 22:Atom 23:Atom 24:Atom 25:Atom 26:Atom 27:Atom 28:Atom 29:CLASS 31:CLASS 32:CLASS 33:CLASS 36:CLASS

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MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),
AND CURRENT DISCOVER FILE IS DATED 11 AUGUST 2004
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=>

L1 STRUCTURE UPLOADED

=> s l1

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SAMPLE SCREEN SEARCH COMPLETED - 1 TO ITERATE

100.0% PROCESSED

1 ITERATIONS

0 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**

PROJECTED ITERATIONS:

1 TO 80

PROJECTED ANSWERS:

0 TO

L2 0 SEA SSS SAM L1

=> s l1 full

THE ESTIMATED SEARCH COST FOR FILE 'REGISTRY' IS 155.00 U.S. DOLLARS DO YOU WANT TO CONTINUE WITH THIS REQUEST? (Y)/N or END:Y
FULL SEARCH INITIATED 12:36:31 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 75 TO ITERATE

100.0% PROCESSED

75 ITERATIONS

0 ANSWERS

SEARCH TIME: 00.00.01

L3 0 SEA SSS FUL L1

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L4 STRUCTURE UPLOADED

=> a 14

L4 HAS NO ANSWERS

L4

STR

=> s 14

SAMPLE SEARCH INITIATED 12:41:47 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 1 TO ITERATE

100.0% PROCESSED

1 ITERATIONS

1 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH

COMPLETE

PROJECTED ITERATIONS:

1 TO 80

PROJECTED ANSWERS:

1 TO 80

L5

1 SEA SSS SAM L4

=> s 14 full

THE ESTIMATED SEARCH COST FOR FILE 'REGISTRY' IS 155.00 U.S. DOLLARS DO YOU WANT TO CONTINUE WITH THIS REQUEST? (Y)/N or END: Y
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h ebc gcg b c

100.0% PROCESSED

4 ITERATIONS

1 ANSWERS

SEARCH TIME: 00.00.02

L6

1 SEA SSS FUL L4

=> file hcaplus

COST IN U.S. DOLLARS

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TOTAL

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316.72

316.93

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FILE COVERS 1907 - 16 Dec 2004 VOL 141 ISS 25 FILE LAST UPDATED: 15 Dec 2004 (20041215/ED)

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3 L6

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ANSWER 1 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN



ACCESSION NUMBER:

2003:719300 HCAPLUS

DOCUMENT NUMBER:

139:240389

TITLE:

Antidepressant

INVENTOR(S): PATENT ASSIGNEE(S): Ohkawa, Shigenori; Miyamoto, Masaomi Takeda Chemical Industries, Ltd., Japan

SOURCE:

PCT Int. Appl., 95 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

Japanese

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PA	TENT	NO.			KIN	D	DATE			APPL	ICAT	ION :	NO.		D	ATE -	
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WO	2003	0740	46		A1		2003	0912		WO 2	003-	JP22	93		2	0030	228
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		GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KR,	ΚZ,	LC,	LK,	LR,	LS,
		LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NO,	NZ,	OM,	PH.	PL.

PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG JP 2004083556 **A**2 20040318 JP 2003-52503 20030228 EP 1481679 EP 2003-707169 **A**1 20041201 20030228 AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK PRIORITY APPLN. INFO.: JP 2002-55771 A 20020301 JP 2002-195434 A 20020704 WO 2003-JP2293 W 20030228 OTHER SOURCE(S): MARPAT 139:240389

GΙ

$$\mathbb{N} = \mathbb{N} - \mathbb{N} - \mathbb{N}$$

AΒ A PKB (Akt) activator contg. a compd. represented by the formula (I) [wherein R1 and R2 each represents hydrogen, a hydrocarbon group, or a heterocyclic group or R1 and R2 form a ring in cooperation with the adjacent carbon atom; R3 represents hydrogen, a hydrocarbon group, or a heterocyclic group; W represents a group represented by the formula (II) (-N(R4)(R5)) or (-XR4c) (wherein ring A represents an optionally substituted benzene ring; ring B represents an optionally substituted 5to 7-membered nitrogenous heterocycle; R4 represents either an arom.-group-substituted aliph. hydrocarbon group which may have other substituent(s) or an acyl contg. an arom. group; R5 represents hydrogen, C1-6 alkyl, or acyl; R4c represents an arom. group, aliph. hydrocarbon group, or acyl; and X represents oxygen or sulfur); Y represents oxygen, sulfur, or NH; and ring C represents an optionally substituted benzene ring], a salt of the compd., or a prodrug of either. Also provided is a use of the activator in or as a preventive/therapeutic agent for depressive psychoses, anxiety disorders, affective psychoses, or PTSD.

IT 216989-41-2

h

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(benzofuran analogs as protein kinase B activators and antidepressants)

RN <u>216989-41-2</u> HCAPLUS

CN Benzofuran, 5-[(4-methoxyphenyl)methoxy]-2,4,6,7-tetramethyl-3-[4-(1methylethyl)phenyl]- (9CI) (CA INDEX NAME)

3 REFERENCE COUNT: THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS

eb c g cg b

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L7 ANSWER 2 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN

FUIL CARREST CO.

ACCESSION NUMBER:

2002:275980 HCAPLUS

DOCUMENT NUMBER:

136:309840

TITLE:

Preparation of heterocyclic compounds as promoters for

the proliferation and differentiation of stem cells

and neuron precursor cells

INVENTOR (S):

Okawa, Shigenori; Miyamoto, Masaomi; Okura, Masahiro

Takeda Chemical Industries, Ltd., Japan

SOURCE:

PCT Int. Appl., 182 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

Japanese

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT ASSIGNEE(S):

	PATENT NO.					KIND DATE			APPLICATION NO.									
	WO 2002028850					A1 20020413												
	w:	ΑE,	AG,	AL,	AM,	AT,	ΑU,	AZ,	BA,	BB,	BG,	BR,	BY,	BZ,	CA,	CH,	CN,	
		CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,	
		GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KR,	ΚZ,	LC,	LK,	LR,	LS,	
		LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NO,	NZ,	PH,	PL,	PT,	
		RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,	ТJ,	TM,	TR,	TT,	TZ,	UA,	UG,	US,	
		UZ,	VN,	YU,	ZA,	ZW,	AM,	AZ,	BY,	KG,	ΚZ,	MD,	RU,	ТJ,	TM			
	RW:	GH,	GM,	KE,	LS,	MW,	ΜZ,	SD,	SL,	SZ,	TZ,	UG,	ZW,	AT,	BE,	CH,	CY,	
		DE,	DK,	ES,	FI,	FR,	GB,	GR,	IE,	IT,	LU,	MC,	NL,	PT,	SE,	TR,	BF,	
		ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,	TD,	ΤG		
	AU 2001092350					A5 20020415				AU 2	001-	9235						
	JP 2002348239					A2 20021204				JP 2	001-	3085	20011004					
	CA 2424870									CA 2	001-	2424	870					
	EP 1323716					A1 20030702				EP 2	001-	9726	87	20011004				
	R:	ΑT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	ΝL,	SE,	MC,	PT,	
		ΙE,	SI,	LT,	LV,	FI,	RO,	MK,	CY,	AL,	TR							
	US 2004034049						A1 20040219			US 2003-398278					20030401			
PRIORITY APPLN. INFO.:										JP 2	000-	3068	01	Ī	A 2	0001	005	
										WO 2	001-	JP87:	39	Ţ	w 2	0011	004	
OTHER SOURCE(S): GI						TAP	136:	30984	40									

$$\mathbb{A} = \mathbb{R}^3$$

AB The title compds. I [R1 and R2 are each H, a hydrocarbon group, a heterocyclic group, or R1 and R2 together with the carbon atom adjacent thereto may form a ring; R3 is H, a hydrocarbon group, or a heterocyclic group; W is R4R5N, etc.; R4 is acyl which is substituted with an arom. group and addnl. bears an optionally substituted aliph. hydrocarbon group or an arom. group; R5 is H, C1-6 alkyl, or acyl; Y is O, S, or NH; and

ring C is an optionally substituted benzene ring] are prepd. Three compds. of this invention at 1 μM gave 344% to 478% promotion of neuron generation. Formulations are given.

IT 216989-41-2P

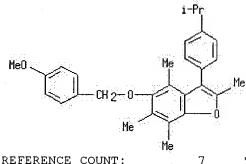
CN

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of heterocyclic compds. as promoters for proliferation and differentiation of stem cells and neuron precursor cells)

RN 216989-41-2 HCAPLUS

Benzofuran, 5-[(4-methoxyphenyl)methoxy]-2,4,6,7-tetramethyl-3-[4-(1-methoxyphenyl)methoxy]-2,4,6,7-tetramethyl-3-[4-(1-methoxyphenyl)methoxyphenyl)methoxyphenylmethylethyl)phenyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT:

THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 3 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER:

1998:806634 HCAPLUS

DOCUMENT NUMBER:

130:38285

TITLE:

Benzofuran derivatives useful for suppressing

neurodegeneration.

INVENTOR(S):

Ohkawa, Shigenori; Setoh, Masaki; Kakihana, Mitsuru;

Okura, Masahiro

PATENT ASSIGNEE(S):

Takeda Chemical Industries, Ltd., Japan

SOURCE:

PCT Int. Appl., 132 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

h

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.					KIND DATE				APPLICATION NO.							DATE		
<u>WO 9855454</u>				A2	A2 19981210				WO 1	998-	19980604							
<u>WO 9855454</u>				А3		1999	0304											
	w:	AL,	AM,	AU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	CA,	CN,	CU,	CZ,	EE,	GE,	GW,	
		HU,	ID,	IL,	IS,	KG,	KR,	KZ,	LC,	LK,	LR,	LT,	LV,	MD,	MG,	MK,	MN,	
		MX,	NO,	ΝZ,	PL,	RO,	RU,	SG,	SI,	SK,	SL,	ТJ,	TM,	TR,	TT,	UA,	US,	
		UZ,	VN,	YU,	AM,	ΑZ,	BY,	KG,	ΚZ,	MD,	RU,	ТJ,	$\mathbf{M}\mathbf{T}$					
	RW:	GH,	GM,	KE,	LS,	MW,	SD,	SZ,	UG,	ΖW,	AT,	BE,	CH,	CY,	DE,	DK,	ES,	
		FI,	FR,	GB,	GR,	IE,	IT,	LU,	MC,	NL,	PT,	SE,	BF,	ВJ,	CF,	CG,	CI,	
		CM,	GA,	GN,	ML,	MR,	ΝE,	SN,	TD,	TG								
AU 9875503				A1		1998	1221		AU 1	998-	19980604							
JP 11049765			A2		1999	0223	1	JP 1:	998-	19980604								
EP 988289			A2		2000	0329	į	EP 1:	998-	19980604								
	R:	AT,	BE.	CH.	DE.	DK.	ES.	FR.	GB.	GR.	IT.	LT.	T.U.	NI.	SE.	MC.	PT.	

IE, FI

PRIORITY APPLN. INFO.:

JP 1997-148325 WO 1998-JP2482

П

19970605 Α 19980604

OTHER SOURCE(S):

MARPAT 130:38285

GT

AΒ Title compds. I [R1, R2 = H, (un)substituted hydrocarbon group; or R1 and R2 form a 3- to 8-membered carbo- or heterocyclic ring which may be substituted; R3 = H, (un)substituted lower alkyl or arom. group; R4 = (un) substituted arom. or araliph. group, or acyl; X , Y = 0 or S which may be oxidized; benzene ring may be further substituted] and their salts are disclosed. The compds. suppress β -amyloid toxicity, and are thus useful as agents for treating of preventing neurodegenerative diseases such as Alzheimer's disease or Parkinsonism. Prepns. of 33 compds. I and their intermediates are described. -- For instance, etherification of 3-(4-isopropylphenyl)-2,2,4,6,7-pentamethyl-2,3-dihydrobenzofuran-5-ol with 4-methoxybenzyl chloride using NaH in DMF gave 49% title compd. II. Seven example compds. gave 27.3-47.0% in vitro protection of human neuroblastoma SK-N-SH cells from β -amyloid neurotoxicity.

IT <u>216989-41-2</u>P, 3-(4-Isopropylphenyl)-5-[(4-methoxybenzyl)oxy]-

2,4,6,7-tetramethylbenzofuran

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(product; prepn. of benzofuran derivs. as agents for suppressing neurodegeneration)

216989-41-2 HCAPLUS RN

Benzofuran, 5-[(4-methoxyphenyl)methoxy]-2,4,6,7-tetramethyl-3-[4-(1-methoxyphenyl)methoxy]-2,4,6,7-tetramethyl-3-[4-(1-methoxyphenyl)methoxy]-2,4,6,7-tetramethyl-3-[4-(1-methoxyphenyl)methoxyphenyl)methoxyl-3-[4-(1-methoxyphenyl)methoxyphenyl)methoxyl-3-[4-(1-methoxyphenyl)meCN methylethyl)phenyl]- (9CI) (CA INDEX NAME)

=> file caold COST IN U.S. DOLLARS

SINCE FILE

TOTAL

FULL ESTIMATED COST ENTRY SESSION 16.64 333.57

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION

-2.10

-2.10

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L6 1 S L4 FULL

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